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In the Claims

Claims 1-30 are pending in this Application.

Original Claims 1-30 are canceled.

Claims 31-58 are new.

31. (new) A compound according to formula I,

or a pharmaceutically acceptable salt or a stereoisomer, thereof, wherein,

A is a five- to ten-membered ring containing up to three heteroatoms; provided A is not a saturated alicyclic when X^2 is =N-, X^3 is -O-, and A is a pyridin-4-yl;

R¹ is selected from -H, halo, trihalomethyl, -CN, -NO₂, -OR⁴, -N(R⁴)R⁴, -S(O)₀₋₂R⁴, $-SO_2N(R^4)R^4$, $-CO_2R^4$, $-C(=O)N(R^4)R^4$, $-C(=O)R^4$, $-C(=NR^5)N(R^4)R^4$, $-C(=NR^5)R^4$, $-N(R^4)SO_2R^4$, $-N(R^4)C(O)R^4$, alkoxy, C_{1-6} alkyl, aryl, aryl, aryl C_{1-6} alkyl, heterocyclyl, and heterocyclyl C₁₋₆ alkyl;

two adjacent of R¹, together with the annular atoms to which they are attached, can form a five- to six-membered ring containing up to two heteroatoms and optionally substituted with up to four of R¹⁰;

R² and R³, together with the annular atoms to which they are attached, form a five- to sixmembered ring containing up to two heteroatoms and optionally substituted with up to five of R⁶:

each R^4 is selected from -H; C_{1-6} alkyl optionally substituted with 1, 2, or 3 halogen; C_{1-6} alkyl optionally substituted with alkoxy; C_{1-6} alkyl substituted with amino where the amino is optionally substituted with one or groups selected from methyl, ethyl, -CH₂CH₂OCH₃, -CH₂CH₂N(CH₃)₂, -CH₂CH₂CH₂N(CH₃)₂, and N-methyl-pyrrolidin3-yl; aryl; aryl C₁₋₆ alkyl; heterocyclyl; and heterocyclyl C₁₋₆ alkyl where the heterocyclyl is

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optionally substituted with alkyl, acyl, NH₂, alkylamino, dialkylamino, heterocyclyl, cyclohexyl, -CH₂OCH₃, -CH₂C(O)NHCH(CH₃)₂, or -CH₂OCH₃;

two of R⁴, when taken together with a common nitrogen to which they are attached, form an five- to seven-membered heterocyclyl, said optionally substituted five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

each R^5 is selected from -H, -CN, -NO₂, -OR⁴, -S(O)₀₋₂R⁴, -CO₂R⁴, C₁₋₆ alkyl, C₂₋₆ alkenyl, and C₂₋₆ alkynyl;

Y is =N- or =
$$C(R^8)$$
-;

 X^1 and X^2 are each independently either =N- or =C(\mathbb{R}^9)-;

 X^3 is selected from $-N(R^7)$ -, -O-, and -S-;

R⁷ is hydrogen;

each of R^6 , R^8 , and R^{10} is independently selected from -H, halo, trihalomethyl, -CN, -NO₂, -OR⁴, -N(R⁴)R⁴, -S(O)₀₋₂R⁴, -SO₂N(R⁴)R⁴, -CO₂R⁴, -C(=O)N(R⁴)R⁴, -C(=NR⁵)N(R⁴)R⁴, -C(=NR⁵)R⁴, -N(R⁴)SO₂R⁴, -N(R⁴)C(O)R⁴, -C(=O)R⁴, optionally substituted alkoxy, C_{1-6} alkyl, aryl, aryl C_{1-6} alkyl, heterocyclyl, and heterocyclyl C_{1-6} alkyl;

two adjacent of R⁶, together with the annular atoms to which they are attached, can form a five- to seven-membered ring containing up to two heteroatoms; and

each R^9 is independently selected from -H; halo; trihalomethyl; -CN; $-NO_2$; $-OR^4$; $-N(R^4)R^4$; $-S(O)_{0-2}R^4$; $-SO_2N(R^4)R^4$; $-CO_2R^4$; $-C(=O)N(R^4)R^4$; $-C(=NR^5)N(R^4)R^4$; $-C(=NR^5)R^4$; $-N(R^4)SO_2R^4$; $-N(R^4)C(O)R^4$; $-C(=O)R^4$; alkoxy; C_{1-6} alkyl optionally substituted with one group selected from alkoxy, benzylamino, and 2-oxo-pyrrolidinyl; aryl C_{1-6} alkyl substituted on the aryl with 1 or 2 groups selected from alkyl and alkoxy; heterocyclyl optionally substituted with -C(O)Ot-Bu; and heterocyclyl C_{1-6} alkyl; provided when R^9 is aryl, heteroaryl, -C(H)=C(H)R or -C(H)=NR, where R is an

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optionally substituted alkyl, cycloalkyl, heteroalicyclic, aryl, or heteroaryl, then Y is not =C(H)-.

- 32. (new) The compound according to claim 31, wherein the five- to six-membered ring formed by R^2 and R^3 is an aryl or a heteroaryl optionally substituted with up to five of R^6 ; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 33. (new) The compound according to claim 32, wherein the five- to six-membered ring formed by R^2 and R^3 is phenyl or pyridyl optionally substituted with up to five of R^6 ; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 34. (new) The compound according to claim 33, of formula II,

$$1-4(R^6)$$
 X^1
 X^2
 X^3

or a pharmaceutically acceptable salt or stereoisomer, thereof.

- 35. (new) The compound according to claim 34, wherein X^1 is $=C(R^9)$ -, X^2 is =N-, X^3 is $-N(R^7)$ -, and R^7 is hydrogen; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 36. (new) The compound according to claim 35, wherein Y is =N-; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 37. (new) The compound according to claim 36, wherein A is either a six- to tenmembered aryl or a five- to ten-membered heteroaryl containing up to three heteroatoms; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 38. (new) The compound according to claim 37, wherein A is either a six-membered aryl or a five- or six-membered heteroaryl containing up to three heteroatoms; or a pharmaceutically acceptable salt or stereoisomer, thereof.

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39. (new) The compound according to claim 38, wherein R^1 is selected from -H, halo, trihalomethyl, -CN, -OR⁴, -N(R⁴)R⁴, -SO₂N(R⁴)R⁴, -CO₂R⁴, -C(=O)N(R⁴)R⁴, -C(=NR⁵)N(R⁴)R⁴, -C(=NR⁵)R⁴, -N(R⁴)SO₂R⁴, -N(R⁴)C(O)R⁴, alkoxy, C₁₋₆ alkyl, heterocyclyl, and heterocyclyl C₁₋₆ alkyl; or a pharmaceutically acceptable salt or stereoisomer, thereof.

40. (new) The compound according to claim 39, of formula III,

wherein R⁷ is hydrogen and at least one of R¹ is -OH; or a pharmaceutically acceptable salt or stereoisomer, thereof.

41. (new) The compound according to claim 40, wherein the compound is either of Formula IIa or IIIb:

$$(R^{6})_{1-4}^{1}$$
 $(R^{6})_{1-4}^{1}$
 $(R^{6})_{1-4}^{1}$
 $(R^{6})_{1-4}^{1}$
 $(R^{1})_{1-4}$
 $(R^{1})_{1-4}$
 $(R^{1})_{1-4}$
 $(R^{1})_{1-4}$

or a pharmaceutically acceptable salt or stereoisomer, thereof.

42. (new) The compound according to claim 41, wherein R^9 is selected from –H; trihalomethyl; C_{1-6} alkyl optionally substituted with one group selected from alkoxy, benzylamino, and 2-oxo-pyrrolidinyl; aryl C_{1-6} alkyl substituted on the aryl with 1 or 2 groups selected from alkyl and alkoxy; heterocyclyl optionally substituted with -C(O)Ot-

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Bu; and heterocyclyl C_{1-6} alkyl; or a pharmaceutically acceptable salt or stereoisomer, thereof.

- 43. (new) The compound according to claim 42, wherein R^6 is selected from -H, halo, trihalomethyl, -CN, -OR⁴, -N(R⁴)R⁴, -CO₂R⁴, -C(=O)N(R⁴)R⁴, -N(R⁴)SO₂R⁴, -N(R⁴)C(O)R⁴, -C(=O)R⁴, C₁₋₆ alkyl, heterocyclyl, heterocyclyl C₁₋₆ alkyl, and a six- or seven-membered heteroalicyclic formed by two adjacent of R^6 , together with the annular atoms to which they are attached, said six- or seven-membered heteroalicyclic containing up to two heteroatoms; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 44. (new) The compound according to claim 43, wherein R^6 is selected from -H, halo, $-OR^4$, $-N(R^4)R^4$, C_{1-6} alkyl, heterocyclyl, heterocyclyl C_{1-6} alkyl, and a six-or sevenmembered heteroalicyclic formed by two adjacent of R^6 , together with the annular atoms to which they are attached, said six- or seven-membered heteroalicyclic containing up to two heteroatoms; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 45. (new) The compound according to claim 44, wherein at least one of R⁶ is -OR⁴ and R⁴ is C₁₋₆ alkyl optionally substituted with 1, 2, or 3 halogen; C₁₋₆ alkyl optionally substituted with alkoxy; C₁₋₆ alkyl substituted with amino where the amino is optionally substituted with one or groups selected from methyl, ethyl, -CH₂CH₂OCH₃, -CH₂CH₂N(CH₃)₂, and *N*-methyl-pyrrolidin3-yl; and heterocyclyl where the heterocyclyl is optionally substituted with alkyl, acyl, NH₂, alkylamino, dialkylamino, heterocyclyl, cyclohexyl, -CH₂OCH₃, -CH₂C(O)NHCH(CH₃)₂, or -CH₂OCH₃; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 46. (new) The compound according to claim 45, wherein at least one of R¹ is halo or methyl; or a pharmaceutically acceptable salt or stereoisomer, thereof.
- 47. (new) The compound according to claim 46, wherein R^9 is selected from -H, trihalomethyl, and C_{1-6} alkyl optionally substituted with one group selected from alkoxy, benzylamino, and 2-oxo-pyrrolidinyl; or a pharmaceutically acceptable salt or stereoisomer, thereof.

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48. (new) The compound according to claim 44, wherein at least one of R^6 is $-OR^4$ and R^4 is heterocyclyl C_{1-6} alkyl where the heterocyclyl is a heteroalicyclic; or a pharmaceutically acceptable salt or stereoisomer, thereof.

The compound according to claim 48, wherein said heteroalicyclic is 49. (new) selected from the group consisting of dioxolanyl, piperidinyl, piperazinyl, 2-oxopiperidinyl, 2-oxopyrrolidinyl, 2-oxoazepinyl, 2-oxopiperazinyl, azepinyl, 4-piperidonyl, pyrrolidinyl, morpholinyl, quinuclidinyl, tetrahydrofuryl, tetrahydropyranyl, thiamorpholinyl, thiamorpholinyl sulfoxide, 2,5-diazabicyclo[2.2.1]heptanyl, and thiamorpholinyl sulfone; or a pharmaceutically acceptable salt or stereoisomer, thereof.

50. (new) The compound according to claim 44, wherein at least one of R⁶ is -OR⁴ and R⁴ is alkyl substituted with at least one additional of alkoxyl, amino, dialkylamino, and monoalkylamino where the monoalkylamino is further sbustituted with *N*-methyl-pyrrolidin3-yl and where each alkyl of monoalkylamino and dialkylamino are independently optionally substituted with -NH₂, -NHCH₃, or -N(CH-)₂; or a pharmaceutically acceptable salt or stereoisomer, thereof.

51. (new) The compound according to claim 31, selected from Table 3; or a pharmaceutically acceptable salt or stereoisomer, thereof

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 10 | 4-{7,8-bis(methyloxy)-1-[(4-methylphenyl)methyl]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | HO—N=NNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNN |

| Entry | Name | Structure |
|-------|--|---|
| 12 | 4-(7,8-bis(methyloxy)-1-{[4-(methyloxy)phenyl]methyl}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | HO N N N N N N N N N N N N N N N N N N N |
| 13 | 4-{7,8-bis(methyloxy)-1-[(2-methylphenyl)methyl]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | HO N N N N |
| .14 | 4-{7,8-bis(methyloxy)-1-[(3-methylphenyl)methyl]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | $HO \longrightarrow N \longrightarrow $ |
| 15 | 4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 20 | 4-[1-{[3,4-bis(methyloxy)phenyl]methyl}-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N = N = N = N = N = N = N = N = N = N = |

| Entry | Name | Structure |
|-------|---|--|
| 21 | 4-(7,8-bis(methyloxy)-1-{[3-(methyloxy)phenyl]methyl}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | HO N N N N N N N N N N N N N N N N N N N |
| 22 | 4-[1-ethyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 25 | 4-[1-methyl-6,7,8- tris(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | HO N N N |
| 27 | 4-[7,8-bis(methyloxy)-1- (trifluoromethyl)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 28 | 4-[1-(1-methylethyl)-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N=NNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNN |

Table 3

| Entry | Name | Structure |
|-------|---|--|
| 29 | 4-[7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N=NN H |
| 31 | 4-[1-methyl-6,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 32 | 4-[6,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 34 | 4-[6,7,8-tris(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 35 | 4-[1-methyl-7,8,9- tris(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | HO N N N N N N N N N N N N N N N N N N N |

| | | able 5 |
|-------|---|---|
| Entry | Name | Structure |
| 36 | 4-[1-methyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 37 | 2-methyl-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 38 | 4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-(methyloxy)phenol | $\begin{array}{c} \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\$ |
| 39 | 4-{1-methyl-8-(methyloxy)-7- [(2-morpholin-4-ylethyl)oxy]- 3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin- 5-yl}phenol | H. Z. |
| 40 | 2-(ethyloxy)-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N |

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Table 3

| Entry | Name | Structure |
|-------|---|--|
| 41 | 2-chloro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N |
| 42 | 2-fluoro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | F N N N N N N N N N N N N N N N N N N N |
| 44 | 2-bromo-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | Br N N H |
| 45 | 1-{[5-(4-hydroxyphenyl)-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-1-yl]methyl}pyrrolidin-2-one | HO N N N |
| 54 | 4-{1-methyl-7-(methyloxy)-8- [(piperidin-4-ylmethyl)oxy]- 3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin- 5-yl}phenol | OH N N N N N N N N N N |

| | 1 | able 3 |
|-------|---|--|
| Entry | Name | Structure |
| 55 | 4-{1-methyl-8-(methyloxy)-7- [(piperidin-4-ylmethyl)oxy]- 3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin- 5-yl}phenol | OH NH |
| 58 | 4-[8-(ethyloxy)-1-methyl-7- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N H |
| 59 | 4-[1-methyl-8,9-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N H |
| 60 | 4-[7-(ethyloxy)-1-methyl-8- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N H |
| 61 | 4-{1-methyl-8-(methyloxy)-9- [(piperidin-4-ylmethyl)oxy]- 3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin- 5-yl}phenol | HO N N N N N N N N N N N N N N N N N N N |

Table 3

| Entry | Name | Structure |
|-------|---|---|
| 63 | 2-ethyl-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 64 | 4-(1-methyl-8-(methyloxy)-9- {[(1-methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | HO NO |
| 65 | 4-(1-methyl-7-(methyloxy)-8- {[(1-methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N, NH OH |
| 66 | 4-[1,7-dimethyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 67 | 1,1-dimethylethyl 4-[5-(4-hydroxyphenyl)-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-1-yl]piperidine-1-carboxylate | HO N N N N N N N N N N N N N N N N N N N |

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 69 | 2-chloro-4-[1-methyl-8,9-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | CI N N N N N N N N N N N N N N N N N N N |
| 70 | 2-fluoro-4-[1-methyl-8,9-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 71 | 2-methyl-4-[1-methyl-8,9-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 72 | 2-bromo-4-[1-methyl-8,9-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | Br O O O O O O O O O O O O O O O O O O O |
| 76 | 2-[(difluoromethyl)oxy]-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | F HO N N N N N |

Table 3

| Entry | Name | Structure |
|-------|---|---|
| 78 | 4-[1,9-dimethyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 79 | 4-[6,9-difluoro-1-methyl-8- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |
| 80 | 2-bromo-4-{1-methyl-8- (methyloxy)-9-[(2-morpholin- 4-ylethyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | N-NH N-NH N-NH N-NH N-NH N-NH N-NH N-NH |
| 81 | 2-chloro-4-{1-methyl-8- (methyloxy)-9-[(2-morpholin- 4-ylethyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | N-NH N-NH O O O O O O O O O O O |
| 82 | 4-(7,8-bis(methyloxy)-1- {[(phenylmethyl)amino]methyl} }-3H-pyrazolo[3,4- c]isoquinolin-5-yl)phenol | $HO \longrightarrow N \longrightarrow $ |

| Entry | Name | Structure |
|-------|--|---|
| 83 | 2,5-dimethyl-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 85 | 2,5-dichloro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | CI N N N N N N N N N N N N N N N N N N N |
| 87 | 2-bromo-4-(1-methyl-8- (methyloxy)-9-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | HO Br O N N N N N N N N N N N N N N N N N N |
| 88 | 2-chloro-4-(1-methyl-7- (methyloxy)-8-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH OH OH |
| 89 | 4-[9-fluoro-1-methyl-8- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- c]isoquinolin-5-yl]phenol | HO — N N N N N N N N N N N N N N N N N N |

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Table 3

| Entry | Name | Structure |
|-------|--|---|
| 90 | 4-(1-methyl-8-(methyloxy)-9- {[2-(methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH NOH OH |
| 91 | 2-chloro-4-(1-methyl-8- (methyloxy)-9-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | HO CI O N N N N N N N N N N N N N N N N N N |
| 92 | 4-[6-bromo-1,7-dimethyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | Br — N — N — N — N — N — N — N — N — N — |
| 93 | 4-[6-fluoro-1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N |
| 94 | 4-[9-chloro-1,7-dimethyl-8- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO CI |

Table 3

| Entry | Name | Structure |
|-------|--|---|
| 95 | 2-chloro-4-[8-{[(1-ethylpiperidin-4-yl)methyl]oxy}-1-methyl-7-(methyloxy)-3 <i>H</i> -pyrazolo[3,4-c]isoquinolin-5-yl]phenol | H Z O O |
| 96 | 3-chloro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO CI N N N N N N N N N N N N N N N N N N |
| 97 | 4-(1-methyl-8,9-bis{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | HO N N N N N N N N N N N N N N N N N N N |
| 98 | 4-(1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | $HO \longrightarrow N \longrightarrow $ |
| 99 | 2-chloro-4-(1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | CI N=N-N |

| Entry | Name | Structure |
|-------|---|--|
| 100 | 2-bromo-4-(1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | HO N N N N N N N N N N N N N N N N N N N |
| 101 | 2-chloro-4-[1,7-dimethyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |
| 102 | 2-bromo-4-[1,7-dimethyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | Br N N N N N N N N N N N N N N N N N N N |
| 103 | 2-chloro-4-[1-methyl-8-({[1-(1-methylethyl)piperidin-4-yl]methyl}oxy)-7-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N-NH OH CI |
| 104 | 4-[9-bromo-1-methyl-8- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO———————————————————————————————————— |

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 105 | 4-[7-chloro-9-fluoro-1-methyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | CI O F |
| 106 | 4-[8-{[(1-acetylpiperidin-4-yl)methyl]oxy}-1-methyl-7-(methyloxy)-3 <i>H</i> -pyrazolo[3,4-c]isoquinolin-5-yl]-2-chlorophenol | N-NH N-NH CI |
| 107 | 4-[9-{[(1-acetylpiperidin-4-yl)methyl]oxy}-1-methyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4-c]isoquinolin-5-yl]-2-bromophenol | By HAN SOLVE O |
| 108 | 2-chloro-4-(1-methyl-9- (methyloxy)-8-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | OH CI |
| 109 | 4-[7-fluoro-1-methyl-8- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N |

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 110 | 2-chloro-4-(1-methyl-8,9-bis{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | |
| 111 - | 2-bromo-4-(1-methyl-8,9-bis{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | Br N N N H |
| 112 | 2-chloro-4-(1-methyl-8- (methyloxy)-9-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH N-OH CI |
| 113 | 2-bromo-4-(1-methyl-8- (methyloxy)-9-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH N O O O O O O O O O Br |

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 114 | 3-fluoro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N H |
| 115 | 2-chloro-4-(1-methyl-7- (methyloxy)-8-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | CI N=N-N H |
| 116 | 2-bromo-4-(1-methyl-7- (methyloxy)-8-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | Br N N N N N N N N N N N N N N N N N N N |
| 117 | 2-chloro-4-(1-methyl-7,8-bis{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | |
| 118 | 2-bromo-4-(1-methyl-7,8-bis{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | Br N N N H |

Table 3

| Entry | Name | Structure |
|-------|--|---|
| 120 | 4-(6,9-difluoro-1-methyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | OH F O O O |
| 121 | 2-chloro-5-fluoro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | CI HO F N H |
| 122 | 2-bromo-5-fluoro-4-[1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | Br N N H |
| 123 | 4-[9-fluoro-1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -benzo[e]indazol-5-yl]phenol | O O O O O O O O O O O O O O O O O O O |
| 125 | 3-fluoro-4-(1-methyl-7- (methyloxy)-8-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | H. Z. |

Table 3

| | Table 5 | | |
|-------|--|--|--|
| Entry | Name | Structure | |
| 126 | 4-(6,9-difluoro-1-methyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)-3-fluorophenol | F F O O F | |
| 127 | 2-chloro-4-(6,9-difluoro-1-methyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | CI F O F | |
| 128 | 2-chloro-4-[8-(ethyloxy)-1-methyl-7-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N N N N N N N N N N N N N N N N N | |
| 129 | 2-chloro-4-[6-chloro-1-methyl-7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | H OH | |
| 130 | 3-fluoro-4-(1-methyl-9- (methyloxy)-8-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | | |

| Entry | Name | Structure |
|-------|--|---|
| 131 | 2-chloro-4-(1,7-dimethyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | CI N N N N N N N N N N N N N N N N N N N |
| 132 | 3-fluoro-4-(6-fluoro-1-methyl-7-(methyloxy)-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | OH F N N N N N N N N N N N N N N N N N N N |
| 133 | 2-chloro-4-[1-methyl-8-[(1-methylethyl)oxy]-7- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | |
| 134 | 2-chloro-4-{1-methyl-7- (methyloxy)-8-[(2- methylpropyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | |
| 135 | 2-bromo-5-fluoro-4-(1-methyl-7-(methyloxy)-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | H N F OH Br |

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Table 3

| Entry | Name | Structure |
|-------|--|--|
| 136 | 4-[7,8-bis(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-chlorophenol | H OH |
| 137 | 4-[7,8-bis(methyloxy)-1- (trifluoromethyl)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]-2-chlorophenol | CI HO N=N N=N N H |
| 138 | 4-{7,8-bis(methyloxy)-1- [(methyloxy)methyl]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}-2-chlorophenol | CI N N N |
| 139 | 2-chloro-4-(1-methyl-3 <i>H</i> -[1,3]dioxolo[4,5-g]pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | HO N N N N N N N N N N N N N N N N N N N |
| 140 | 2-chloro-4-(1-methyl-8,9-dihydro-3 <i>H</i> -[1,4]dioxino[2,3-g]pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | CI N H |

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 141 | 2-chloro-4-(1-methyl-9,10-dihydro-3 <i>H</i> ,8 <i>H</i> -[1,4]dioxepino[2,3-g]pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | CI N N N N N N N N N N N N N N N N N N N |
| 142 | 2-chloro-4-[7- [(difluoromethyl)oxy]-1- methyl-8-(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | F F O O O O O O O O O O O O O O O O O O |
| 143 | 2-chloro-5-fluoro-4-(6-fluoro- 1-methyl-9-(methyloxy)-8-{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH N F OH CI |
| 144 | 2-chloro-4-(6,9-difluoro-1-methyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)-5-fluorophenol | $\begin{array}{c c} CI & F & F \\ HO & N & N \\ F & N & H \end{array}$ |
| 145 | 2-chloro-4-(11-methyl-2,3-dihydro-9 <i>H</i> -[1,4]dioxino[2,3-f]pyrazolo[3,4- <i>c</i>]isoquinolin-7-yl)phenol | CI N N N N N N N N N N N N N N N N N N N |

| Entry | Name | Structure |
|-------|---|--|
| 146 | 2-chloro-5-fluoro-4-(11-methyl-2,3-dihydro-9 <i>H</i> -[1,4]dioxino[2,3-f]pyrazolo[3,4- <i>c</i>]isoquinolin-7-yl)phenol | N-NH N-NH OH CI |
| 147 | 2-chloro-4-[1-methyl-6,7,8- tris(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | CI O N N N N N N N N N N N N N N N N N N |
| 148 | 2-bromo-4-(6,9-difluoro-1-methyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)-5-fluorophenol | Br F N N N H |
| 149 | 7-(3-chlorophenyl)-11-methyl- 2,3-dihydro-9 <i>H</i> - [1,4]dioxino[2,3- f]pyrazolo[3,4- <i>c</i>]isoquinoline | NH NH CI |
| 150 | 2-chloro-5-fluoro-4-(6-fluoro- 1-methyl-8,9-dihydro-3 <i>H</i> - [1,4]dioxino[2,3- g]pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | CI F N N H |

Table 3

| Entry | Name | Structure |
|-------|---|---------------------------|
| 151 | 2-chloro-4-{1-methyl-7- (methyloxy)-8- [(tetrahydrofuran-2- ylmethyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | N-NH N-NH CI |
| 152 | 2-chloro-4-{1-methyl-7- (methyloxy)-8-[(tetrahydro-2 <i>H</i> - pyran-2-ylmethyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | N-NH N-NH CI |
| 153 | 2-chloro-4-{1-methyl-7- (methyloxy)-8-[(2,2,2- trifluoroethyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | F F O CI |
| 154 | 2-chloro-5-fluoro-4-[9-fluoro-1-methyl-6,7,8-tris(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | CI O F HO N N N |
| 155 | 5-(3-chloro-4-hydroxyphenyl)-6,9-difluoro-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | N-NH N F HO F OH |

Table 3

| Entry | Name | Structure |
|-------|---|---------------------------------------|
| 156 | 6,9-difluoro-5-(2-fluorophenyl)-1-methyl-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinoline | N, NH N F N F |
| 157 | 2-chloro-4-{8- [(difluoromethyl)oxy]-6,9- difluoro-1-methyl-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | F F OH CI |
| 158 | 2-chloro-4-(6,11-difluoro-1-methyl-8,9-dihydro-3 <i>H</i> -[1,4]dioxino[2,3-g]pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)-5-fluorophenol | CI F F HO N N N H |
| 159 | 4-(1-methyl-3 <i>H</i> -benzo[e]indazol-5-yl)phenol | N N N N N N N N N N N N N N N N N N N |
| 160 | 6-fluoro-7-(2-fluorophenyl)-11- methyl-2,3-dihydro-9 <i>H</i> - [1,4]dioxino[2,3- f]pyrazolo[3,4- <i>c</i>]isoquinoline | N-NH N F |
| 161 | 2-chloro-4-{1-methyl-7- (methyloxy)-8-[(tetrahydro-2 <i>H</i> - pyran-4-ylmethyl)oxy]-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl}phenol | N-NH N-NH CI |

| | | able 5 |
|-------|--|--|
| Entry | Name | Structure |
| 162 | 2-chloro-4-[8-{[2- (ethyloxy)ethyl]oxy}-1-methyl- 7-(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | CI N N N N N N N N N N N N N N N N N N N |
| 164 | 3-fluoro-4-(6-fluoro-11-methyl- 2,3-dihydro-9 <i>H</i> - [1,4]dioxino[2,3- f]pyrazolo[3,4- <i>c</i>]isoquinolin-7- yl)phenol | N-NH N F O F OH |
| 165 | 2-chloro-5-fluoro-4-(6-fluoro- 11-methyl-2,3-dihydro-9 <i>H</i> - [1,4]dioxino[2,3- f]pyrazolo[3,4- <i>c</i>]isoquinolin-7- yl)phenol | N-NH N F OH CI |
| 166 | 2-chloro-4-[8-(cyclopentyloxy)-1-methyl-7-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N-NH N-NH OH OH |
| 167 | 2-chloro-4-(1-methyl-7-(1-methylethyl)-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | HO N N N |

Table 3

| Entry | Name | Structure |
|-------|---|--|
| 168 | 2-chloro-4-[9-ethyl-1-methyl-8-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | $\begin{array}{c} CI \\ HO \\ \end{array}$ |
| 169 | 2-chloro-4-(6,9-difluoro-1-methyl-8-{[(1-methylpiperidin-4-yl)methyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | N-NH F OH CI |
| 170 | 5-(3-chloro-4-hydroxyphenyl)- 8-fluoro-1-methyl-7- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-6-ol | HO HO N N N N N N N N N N N N N N N N N |
| 171 | 2-chloro-4-(6-fluoro-1-methyl-7-(methyloxy)-8-{[(1-methylpiperidin-4-yl)methyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | NH NH CI |
| 172 | 2-chloro-4-(6-fluoro-1-methyl- 8,9-bis{[2- (methyloxy)ethyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | CI F N N N H |

Table 3

| Entry | Name | Structure |
|-------|---|------------------------------------|
| 173 | 5-[3-chloro-4- (methyloxy)phenyl]-6-fluoro-1- methyl-7-(methyloxy)-8-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinoline | N, NH N CI |
| 174 | 5-[3-chloro-4- (methyloxy)phenyl]-8-fluoro-1- methyl-7-(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-6-ol | HO N N N |
| 176 | 2-chloro-4-{6-fluoro-1-methyl-7-(methyloxy)-8-[(2-piperidin-1-ylethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N-NH O-N-O-H-CI |
| 177 | 2-chloro-4-[8-{[2-(4-ethylpiperazin-1-yl)ethyl]oxy}-6-fluoro-1-methyl-7-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | PH Z OH OH OH |
| 178 | 2-chloro-4-[8-{[2- (diethylamino)ethyl]oxy}-6- fluoro-1-methyl-7-(methyloxy)- 3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin- 5-yl]phenol | N-NH N-NH N-NH N-OH CI |

Table 3

| Entry | Name | Structure |
|-------|---|--|
| 179 | 2-chloro-4-(6-fluoro-1-methyl-7-(methyloxy)-8-{[2-(methyloxy)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | CI F N N N N N N N N N N N N N N N N N N |
| 182 | 2-chloro-4-(6-fluoro-1-methyl- 9-(methyloxy)-8-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH N-NH N-NH N-NH N-OH CI |
| 183 | 2-bromo-4-(6-fluoro-1-methyl- 9-(methyloxy)-8-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | P O O Br |
| 184 | 2-chloro-5-fluoro-4-(6-fluoro-1-methyl-9-(methyloxy)-8-{[(1-methylpiperidin-4-yl)methyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | H Z F CI |
| 185 | 4-(6-fluoro-1-methyl-9- (methyloxy)-8-{[(1- methylpiperidin-4- yl)methyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)-2-methylphenol | N-NH N-NH N-NH N-NH N-NH N-NH N-NH N-NH |

Table 3

| Entry | Name | Structure |
|-------|--|--|
| 186 | 2-chloro-4-{6,9-difluoro-1-methyl-8-[(2-piperidin-1-ylethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N-NH N-NH N-NH N-NH N-NH N-NH N-NH N-NH |
| 187 | 2-chloro-4-(8-{[2-(4-ethylpiperazin-1-yl)ethyl]oxy}-6,9-difluoro-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | E S S S S S S S S S S S S S S S S S S S |
| 188 | 2-chloro-4-(8-{[2- (diethylamino)ethyl]oxy}-6,9- difluoro-1-methyl-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N-NH N-NH N-NH OH CI |
| 191 | 6,9-difluoro-5-(1 <i>H</i> -indol-5-yl)- 1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | HN N N N N N N N N N N N N N N N N N N |
| 193 | 5-(4-aminophenyl)-6,9- difluoro-1-methyl-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | H_2N N N N N N N N |

| Entry | Name | Structure |
|-------|--|--|
| 194 | 2-chloro-4-(6-fluoro-1-methyl-7-(methyloxy)-8-{[2-(4-methylpiperazin-1-yl)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4-c]isoquinolin-5-yl)phenol | N-NH N-NH N-NH N-NH OH CI |
| 195 | 5-(2-amino-1,3-thiazol-5-yl)-6,9-difluoro-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | N-NH N S NH ₂ |
| 196 | 2-chloro-4-[8-{[2-(4-ethylpiperazin-1-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI P CI |
| 197 | 2-chloro-4-(6-fluoro-1-methyl-9-(methyloxy)-8-{[2-(4-methylpiperazin-1-yl)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | H Z OH OH |
| 198 | 5-(6-aminopyridin-3-yl)-6,9-difluoro-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | H_2N N N N N N N N N N |

| Entry | Name | Structure |
|-------|---|--|
| 199 | 5-(5-amino-2-thienyl)-6,9- difluoro-1-methyl-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | H ₂ N S N N N N N N N N N N N N N N N N N N |
| 200 | 2-chloro-4-[8-{[3-(4-ethylpiperazin-1-yl)propyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N NH N CI OH |
| 201 | 2-chloro-4-(6-fluoro-1-methyl- 9-(methyloxy)-8-{[3-(4- methylpiperazin-1- yl)propyl]oxy}-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl)phenol | N, NH N O F O O O O O O O O O O O O O O O O O |
| 202 | 6,9-difluoro-5-(1 <i>H</i> -indol-6-yl)- 1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | F HO F |
| 203 | N-[5-(6,9-difluoro-8-hydroxy-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)-1,3-thiazol-2-yl]acetamide | OH F-F N S N S N S N S N S N S N S N S N S N S |

Table 3

| TR 4 | | St |
|-------|--|--|
| Entry | Name | Structure |
| 206 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(2-morpholin-4-ylethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N-NH N-NH N-NH N-NH N-NH N-NH N-NH N-NH |
| 207 | 4-[8-({2- [butyl(ethyl)amino]ethyl}oxy)- 6-fluoro-1-methyl-9- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2- chlorophenol | N-NH N-NH N-NH N-NH N-NH N-NH N-NH N-NH |
| 208 | 4-[8-{[(2R)-2-amino-3-methylbutyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-chlorophenol | N-NH N-NH N-NH N-NH OH CI |
| 209 | 2-chloro-4-(6-fluoro-1-methyl-9-(methyloxy)-8-{[2-(1-methylpiperidin-4-yl)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | N-NH N-NH N-NH N-OH CI |
| 210 | 2-chloro-4-[8-{[(1-ethylpiperidin-4-yl)methyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N-NH N-NH O-H-CI |

Table 3

| E4 | | Stanisting |
|-------|--|--|
| Entry | Name | Structure |
| 212 | 5-(5-amino-1,3,4-thiadiazol-2-yl)-6,9-difluoro-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | F HO S NH ₂ |
| 213 | 4-[8-{[(2R)-2-amino-3,3-dimethylbutyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-chlorophenol | CI F O NH2 |
| 214 | 2-chloro-4-[6-fluoro-1-methyl-9-(methyloxy)-8-({2-[4-(2-methylpropyl)piperazin-1-yl]ethyl}oxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI F NN NN |
| 215 | 2-chloro-4-[8-{[2-(5-ethyl-2,5-diazabicyclo[2.2.1]hept-2-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyṛazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N O F CI |
| 216 | 2-chloro-4-[6-fluoro-1-methyl-8-({2-[4-(1-methylethyl)piperazin-1-yl]ethyl}oxy)-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI F N N N N N N N N N N N N N N N N N N |

| | Table 3 | | |
|-------|---|--|--|
| Entry | Name | Structure | |
| 217 | 4-[8-{[2-(3-amino-8-azabicyclo[3.2.1]oct-8-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-chlorophenol | H_2N O F O O F O | |
| 218 | 2-chloro-4-[8-{[2-(1-ethylpiperidin-4-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI F N N N N N N N N N N N N N N N N N N | |
| 219 | 2-chloro-4-[8-{[2- (diethylamino)ethyl]oxy}-6- fluoro-1-methyl-9-(methyloxy)- 3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin- 5-yl]phenol | N-NH N-NH O-N-NH | |
| 220 | 2-chloro-5-{6-fluoro-1-methyl-9-(methyloxy)-8-[(2-pyrrolidin-1-ylethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N NH OH CI | |
| 223 | 2-chloro-4-[6-fluoro-1-methyl-8-[(8-methyl-8-azabicyclo[3.2.1]oct-3-yl)oxy]-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI F NNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNNN | |

| Entry | Name | Structure |
|-------|--|--|
| 224 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(3-pyrrolidin-1-ylpropyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N-NH N O F O CI |
| 225 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(3-piperidin-1-ylpropyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | H Z OH OH |
| 226 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(3-morpholin-4-ylpropyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N-NH N-NH N-NH N-NH N-OH CI |
| 227 | 2-chloro-4-[8-({2-[[2-(dimethylamino)ethyl](methyl) amino]ethyl}oxy)-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N H Z O H CI |
| 228 | 2-chloro-4-[8-({2-[[2- (diethylamino)ethyl](methyl)a mino]ethyl}oxy)-6-fluoro-1- methyl-9-(methyloxy)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | OH CI N N N N N N N N N N N N N N N N N N |

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Table 3

| Entry | Name | Structure |
|-------|--|---|
| 229 | 2-chloro-4-[8-({2-[[2-(dimethylamino)ethyl](ethyl)a mino]ethyl}oxy)-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI P N N N N N N N N N N N N N N N N N N |
| 230 | 4-[8-[(2-{bis[3- (dimethylamino)propyl]amino} ethyl)oxy]-6-fluoro-1-methyl-9- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2- chlorophenol | OH CI N N N N N N N N N N N N N N N N N N |
| 231 | 2-chloro-4-[6-fluoro-1-methyl-8-({2-[methyl(1-methylpyrrolidin-3-yl)amino]ethyl}oxy)-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N-NH N O F CI |
| 232 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(2-{(2S)-2-[(methyloxy)methyl]pyrrolidin-1-yl}ethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | HO F O N O N O N O N O N O N O N O N O N |
| 233 | 2-chloro-4-(6-fluoro-1-methyl-9-(methyloxy)-8-{[2-(4-pyrrolidin-1-ylpiperidin-1-yl)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | N-NH N O F O CI |

Table 3

| Entry | Name | Structure |
|-------|---|---|
| 234 | 2-chloro-4-[8-{[2-(4-cyclohexylpiperazin-1-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | H NH N OH CI |
| 235 | 2-[4-(2-{[5-(3-chloro-4-hydroxyphenyl)-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-yl]oxy}ethyl)piperazin-1-yl]- <i>N</i> -(1-methylethyl)acetamide | OH CI PE CO |
| 236 | 4-[8-{[2-(1,4'-bipiperidin-1'-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-chlorophenol | HO N N N N N N N N N N N N N N N N N N N |
| 237 | 2-chloro-4-[6-fluoro-1-methyl-8-{[2-(4-methyl-1,4-diazepan-1-yl)ethyl]oxy}-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | F N N N N N N N N N N N N N N N N N N N |
| 238 | 2-chloro-4-(6-fluoro-1-methyl-9-(methyloxy)-8-{[2-(4-pyridin-2-ylpiperazin-1-yl)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | H Z O O O O O O O O O O O O O O O O O O |

Table 3

| Entry | Name | Structure |
|-------|---|---|
| 239 | 2-chloro-4-[8-{[2-(2,6-dimethylmorpholin-4-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | OH CI F O N N N N N N N N N N N N N N N N N N |
| 240 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(2-thiomorpholin-4-ylethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | S N O F OH |
| 241 | 2-chloro-4-[8-{[2-(2,6-dimethylpiperidin-1-yl)ethyl]oxy}-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | N-NH N-NH O F CI |
| 242 | 2-chloro-4-(6-fluoro-1-methyl-9-(methyloxy)-8-{[2-(octahydroquinolin-1(2 <i>H</i>)-yl)ethyl]oxy}-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl)phenol | N-NH N-NH N-NH OH CI |
| 243 | 4-[8-({2-[bis(1-methylethyl)amino]ethyl}oxy)-6-fluoro-1-methyl-9-(methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2-chlorophenol | N-NH N-NH N-NH OH CI |

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Table 3

| Entry | Name | Structure |
|-------|---|--|
| 244 | 4-[8-[(2-{bis[2- (methyloxy)ethyl]amino}ethyl) oxy]-6-fluoro-1-methyl-9- (methyloxy)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]-2- chlorophenol | OH CI |
| 245 | 2-chloro-4-{6-fluoro-1-methyl-9-(methyloxy)-8-[(2-piperidin-1-ylethyl)oxy]-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl}phenol | N-NH N-NH N-NH N-NH N-NH N-NH N-NH N-NH |

52. (new) A Compound selected from

| 9 | 4-[7,8-bis(methyloxy)-1- (phenylmethyl)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]benzene-1,2-diol | O O O O O O O O O O |
|----|--|---------------------------------------|
| 19 | 4-[7,8-bis(methyloxy)-1-(1-phenylethyl)-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-5-yl]phenol | HO N N N |

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| 24 | 4-[6,7,8-tris(methyloxy)-1- (phenylmethyl)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | N = N = N = N = N = N = N = N = N = N = |
|-----|--|---|
| 26 | 4-[8-(methyloxy)-1- (phenylmethyl)-3 <i>H</i> - pyrazolo[3,4- <i>c</i>]isoquinolin-5- yl]phenol | HO N=N-N |
| 221 | 6,9-difluoro-5-(2-imino-3-methyl-2,3-dihydro-1,3-thiazol-5-yl)-1-methyl-3 <i>H</i> -pyrazolo[3,4- <i>c</i>]isoquinolin-8-ol | H S NH F HO |

- 53. (new) A pharmaceutical composition comprising the compound according to claim 31 and a pharmaceutically acceptable carrier.
- 54. (new) A method of modulating the *in vivo* activity of a kinase, the method comprising administering to a subject an effective amount of the compound or the pharmaceutical composition according to claim 31.
- 55. (new) The method according to claim 54, wherein the kinase is ALK.
- 56. (new) The method according to claim 55, wherein modulating the *in vivo* activity of ALK comprises inhibition of ALK.
- 57. (new) A method of treating diseases or disorders associated with uncontrolled, abnormal, and/or unwanted cellular activities, the method comprising administering, to a

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mammal in need thereof, a therapeutically effective amount of the compound or the pharmaceutical composition as described in any one of claims 31.

The method of claim 57 where the disease is an ALK-positive lymphomas, 58. (new) B-cell lymphoma, neuroblastoma, or inflammatory myofibroblastic tumors.